

**WHAT IS CLAIMED IS:**

- 1           1. A method for identifying a compound that modulates cell cycle  
2 arrest, the method comprising the steps of:
  - 3                 (i) contacting a cell comprising a target polypeptide or fragment thereof or  
4 inactive variant thereof, selected from the group consisting of flap structure specific  
5 endonuclease 1 (FEN1), protein kinase C  $\zeta$  (PKC- $\zeta$ ), phospholipase C- $\beta$ 1 (PLC- $\beta$ 1),  
6 protein tyrosine kinase 2 (FAK), protein tyrosine kinase 2b (FAK2), casein kinase 2  
7 (CK2), cMET tyrosine kinase (cMET), REV1 dCMP transferase (REV1),  
8 apurinic/apyrimidinic nuclease 1 (APE1), cyclin dependent kinase 3 (CDK3), PIM1  
9 kinase (PIM1), cell division cycle 7 kinase (CDC7L1), cyclin dependent kinase 7  
10 (CDK7), cytokine inducible kinase (CNK), potentially prenylated protein tyrosine  
11 phosphatase (PRL-3), serine threonine kinase 2 (STK2) or (NEK4), cyclin dependent  
12 serine threonine kinase (NKIAMRE), or histone acetylase (HBO1), or fragment thereof  
13 with the compound, the target polypeptide encoded by the complement of a nucleic acid  
14 that hybridizes under stringent conditions to a nucleic acid encoding a polypeptide having  
15 an amino acid sequence selected from the group consisting of SEQ ID NO:14, 2, 4, 6, 8,  
16 10, 12, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, and 36; and
  - 17                 (ii) determining the chemical or phenotypic effect of the compound upon  
18 the cell comprising the target polypeptide or fragment thereof or inactive variant thereof,  
19 thereby identifying a compound that modulates cell cycle arrest.
- 1           2. The method of claim 1, wherein the chemical or phenotypic effect  
2 is determined by measuring enzymatic activity selected from the group consisting of  
3 nuclease activity, kinase activity, lipase activity, transferase activity, phosphatase activity,  
4 and acetylase activity.
- 1           3. The method of claim 1, wherein the chemical or phenotypic effect  
2 is determined by measuring cellular proliferation.
- 1           4. The method of claim 3, wherein the cellular proliferation is  
2 measured by assaying fluorescent marker level or DNA synthesis.
- 1           5. The method of claim 4, wherein DNA synthesis is measured by  $^3\text{H}$   
2 thymidine incorporation, BrdU incorporation, or Hoescht staining.

1                   6.     The method of claim 4, wherein the fluorescent marker is selected  
2 from the group consisting of a cell tracker dye or green fluorescent protein.

1                   7.     The method of claim 1, wherein modulation is activation of cell  
2 cycle arrest.

1                   8.     The method of claim 1, wherein modulation is activation of cancer  
2 cell cycle arrest.

1                   9.     The method of claim 1, wherein the host cell is a cancer cell.

1                   10.    The method of claim 9, wherein the cancer cell is a breast, prostate,  
2 colon, or lung cancer cell.

1                   11.    The method of claim 9, wherein the cancer cell is a transformed  
2 cell line.

1                   12.    The method of claim 11, wherein the transformed cell line is A549,  
2 PC3, H1299, MDA-MB-231, MCF7, or HeLa.

1                   13.    The method of claim 9, wherein the cancer cell is p53 null or  
2 mutant.

1                   14.    The method of claim 9, wherein the cancer cell is p53 wild-type.

1                   15.    The method of claim 1, wherein the polypeptide is recombinant.

1                   16.    The method of claim 1, wherein the polypeptide is encoded by a  
2 nucleic acid comprising a sequence of SEQ ID NO:13, 1, 3, 5, 7, 9, 11, 15, 17, 19, 21, 23,  
3 25, 27, 29, 31, 33, or 35.

1                   17.    The method of claim 1, wherein the compound is an antibody.

1                   18.    The method of claim 1, wherein the compound is a small organic  
2 molecule.

1                   19.    The method of claim 1, wherein the compound is an antisense  
2 molecule.

1                   20. The method of claim 1, wherein the compound is a peptide.

1                   21. The method of claim 20, wherein the peptide is circular.

1                   22. The method of claim 1, wherein the compound is an siRNA  
2 molecule.

1                   23. A method for identifying a compound that modulates cell cycle  
2 arrest, the method comprising the steps of:

3                   (i) contacting a cell comprising a target polypeptide or fragment thereof or  
4 inactive variant thereof, selected from the group consisting of flap structure specific  
5 endonuclease 1 (FEN1), protein kinase C  $\zeta$  (PKC- $\zeta$ ), phospholipase C- $\beta$ 1 (PLC- $\beta$ 1),  
6 protein tyrosine kinase 2 (FAK), protein tyrosine kinase 2b (FAK2), casein kinase 2  
7 (CK2), cMET tyrosine kinase (cMET), REV1 dCMP transferase (REV1),  
8 apurinic/apyrimidinic nuclelease 1 (APE1), cyclin dependent kinase 3 (CDK3), PIM1  
9 kinase (PIM1), cell division cycle 7 kinase (CDC7L1), cyclin dependent kinase 7  
10 (CDK7), cytokine inducible kinase (CNK), potentially prenylated protein tyrosine  
11 phosphatase (PRL-3), serine threonine kinase 2 (STK2) or (NEK4), cyclin dependent  
12 serine threonine kinase (NKIAMRE), or histone acetylase (HBO1), or fragment thereof  
13 with the compound, the target polypeptide encoded by the complement of a nucleic acid  
14 that hybridizes under stringent conditions to a nucleic acid encoding a polypeptide having  
15 an amino acid sequence selected from the group consisting of SEQ ID NO:14, 2, 4, 6, 8,  
16 10, 12, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, and 36; and

17                   (ii) determining the physical effect of the compound upon the target  
18 polypeptide or fragment thereof or inactive variant thereof; and

19                   (iii) determining the chemical or phenotypic effect of the compound upon  
20 a cell comprising the target polypeptide or or fragment thereof or inactive variant thereof,  
21 thereby identifying a compound that modulates cell cycle arrest.

1                   24. A method of modulating cell cycle arrest in a subject, the method  
2 comprising the step of administering to the subject a therapeutically effective amount of a  
3 compound identified using the method of claim 1.

1                   25. The method of claim 24, wherein the subject is a human.

- 1                   26. The method of claim 25, wherein the subject has cancer.
- 1                   27. The method of claim 24, wherein the compound is a small organic  
2 molecule.
- 1                   28. The method of claim 24, wherein the compound is an antisense  
2 molecule.
- 1                   29. The method of claim 24, wherein the compound is an antibody.
- 1                   30. The method of claim 24, wherein the compound is a peptide.
- 1                   31. The method of claim 30, wherein the peptide is circular.
- 1                   32. The method of claim 24, wherein the compound is an siRNA  
2 molecule.
- 1                   33. The method of claim 24, wherein the compound inhibits cancer cell  
2 proliferation.
- 1                   34. A method of modulating cell cycle arrests in a subject, the method  
2 comprising the step of administering to the subject a therapeutically effective amount of a  
3 target polypeptide or fragment thereof or inactive variant thereof, selected from the group  
4 consisting of flap structure specific endonuclease 1 (FEN1), protein kinase C  $\zeta$  (PKC- $\zeta$ ),  
5 phospholipase C- $\beta$ 1 (PLC- $\beta$ 1), protein tyrosine kinase 2 (FAK), protein tyrosine kinase  
6 2b (FAK2), casein kinase 2 (CK2), cMET tyrosine kinase (cMET), REV1 dCMP  
7 transferase (REV1), apurinic/apyrimidinic nuclease 1 (APE1), cyclin dependent kinase 3  
8 (CDK3), PIM1 kinase (PIM1), cell division cycle 7 kinase (CDC7L1), cyclin dependent  
9 kinase 7 (CDK7), cytokine inducible kinase (CNK), potentially prenylated protein  
10 tyrosine phosphatase (PRL-3), serine threonine kinase 2 (STK2) or (NEK4), cyclin  
11 dependent serine threonine kinase (NKIAMRE), or histone acetylase (HBO1), or  
12 fragment thereof with the compound, the target polypeptide encoded by the complement  
13 of a nucleic acid that hybridizes under stringent conditions to a nucleic acid encoding a  
14 polypeptide having an amino acid sequence selected from the group consisting of SEQ ID  
15 NO:14, 2, 4, 6, 8, 10, 12, 16, 18, 20, 22, 24, 26, 28, 30, 32, 34, and 36.

1               35.     A method of modulating cell cycle arrest in a subject, the method  
2     comprising the step of administering to the subject a therapeutically effective amount of a  
3     nucleic acid encoding a target polypeptide or fragment thereof or inactive variant thereof,  
4     selected from the group consisting of flap structure specific endonuclease 1 (FEN1), protein  
5     kinase C  $\zeta$  (PKC- $\zeta$ ), phospholipase C- $\beta$ 1 (PLC- $\beta$ 1), protein tyrosine kinase 2 (FAK), protein  
6     tyrosine kinase 2b (FAK2), casein kinase 2 (CK2), cMET tyrosine kinase (cMET), REV1  
7     dCMP transferase (REV1), apurinic/apyrimidinic nuclelease 1 (APE1), cyclin dependent  
8     kinase 3 (CDK3), PIM1 kinase (PIM1), cell division cycle 7 kinase (CDC7L1), cyclin  
9     dependent kinase 7 (CDK7), cytokine inducible kinase (CNK), potentially prenylated protein  
10    tyrosine phosphatase (PRL-3), serine threonine kinase 2 (STK2) or (NEK4), cyclin dependent  
11    serine threonine kinase (NKIAMRE), or histone acetylase (HBO1), or fragment thereof with  
12    the compound, the target polypeptide encoded by the complement of a nucleic acid that  
13    hybridizes under stringent conditions to a nucleic acid encoding a polypeptide having an  
14    amino acid sequence selected from the group consisting of SEQ ID NO:14, 2, 4, 6, 8, 10, 12,  
15    16, 18, 20, 22, 24, 26, 28, 30, 32, 34, and 36.

1               36.     A CK2-specific siRNA molecule comprising the sequence  
2     AACATTGAATTAGATCCACGT, wherein the siRNA molecule is from 21 to 30 nucleotide  
3     base pairs in length.

1               37.     The CK2-specific siRNA molecule of claim 36 consisting of the  
2     sequence AACATTGAATTAGATCCACGT and its complement as active portion.

1               38.     A method of inhibiting expression of a CK2 gene in a cell, the method  
2     comprising contacting the cell with a CK2-specific siRNA molecule comprising the sequence  
3     AACATTGAATTAGATCCACGT, wherein the siRNA molecule is from 21 to 30 nucleotide  
4     base pairs in length.

1               39.     A PIM1-specific siRNA molecule comprising the sequence  
2     AAAACCTCCGAGTGAACGGTC, wherein the siRNA molecule is from 21 to 30  
3     nucleotide base pairs in length.

1               40.     The PIM1-specific siRNA molecule of claim 39 consisting of the  
2     sequence AAAACCTCCGAGTGAACGGTC and its complement as active portion.

1                  41.     A method of inhibiting expression of a PIM1 gene in a cell, the method  
2   comprising contacting the cell with a PIM1-specific siRNA molecule comprising the  
3   sequence AAAACTCCGAGTGAACGGTC, wherein the siRNA molecule is from 21 to 30  
4   nucleotide base pairs in length.

1                  42.     An Hbo1-specific siRNA molecule comprising the sequence  
2   AACTGAGCAAGTGGTTGATT, wherein the siRNA molecule is from 21 to 30 nucleotide  
3   base pairs in length.

1                  43.     The Hbo1-specific siRNA molecule of claim 42 consisting of the  
2   sequence AACTGAGCAAGTGGTTGATT and its complement as active portion.

1                  44.     A method of inhibiting expression of an Hbo1 gene in a cell, the  
2   method comprising contacting the cell with an Hbo1-specific siRNA molecule comprising  
3   the sequence AACTGAGCAAGTGGTTGATT, wherein the siRNA molecule is from 21 to  
4   30 nucleotide base pairs in length.